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(54) **HYDROGEL COMPOSITES AND SUPERPOROUS HYDROGEL COMPOSITES HAVING FAST SWELLING, HIGH MECHANICAL STRENGTH, AND SUPERABSORBENT PROPERTIES**

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(58) Field of Search ..... **521/102, 109.1, 521/121, 125, 128, 130, 140, 142, 146, 149, 150, 182, 183, 186, 187**

(56) **References Cited**

#### U.S. PATENT DOCUMENTS

|           |           |                       |         |
|-----------|-----------|-----------------------|---------|
| 3,551,556 | 12/1970   | Kliment et al. .      |         |
| 3,641,237 | 2/1972    | Gould et al. .        |         |
| 3,826,678 | 7/1974    | Hoffman et al. .      |         |
| 4,178,361 | 12/1979   | Cohen et al. .        |         |
| 4,525,527 | 6/1985    | Takeda et al. .       |         |
| 4,529,739 | 7/1985    | Scott et al. .        |         |
| 4,649,164 | 3/1987    | Scott et al. .        |         |
| 4,801,457 | 1/1989    | Heller et al. .       |         |
| 5,002,814 | 3/1991    | Knack et al. .        |         |
| 5,089,606 | 2/1992    | Cole et al. .         |         |
| 5,147,343 | 9/1992    | Kellenberger .        |         |
| 5,149,335 | 9/1992    | Kellenberger et al. . |         |
| 5,154,713 | 10/1992   | Lind .                |         |
| 5,292,777 | 3/1994    | DesMarais et al. .    |         |
| 5,324,561 | 6/1994    | Rezai et al. .        |         |
| 5,338,766 | 8/1994    | Phan et al. .         |         |
| 5,352,448 | 10/1994   | Bowerstock et al. .   |         |
| 5,403,870 | 4/1995    | Gross .               |         |
| 5,424,265 | * 6/1995  | Weinstein             | 502/400 |
| 5,451,613 | 9/1995    | Smith et al. .        |         |
| 5,462,972 | * 10/1995 | Smith et al. .        | 521/64  |
| 5,624,967 | * 4/1997  | Hitomi et al. .       | 521/64  |
| 5,750,585 | 5/1998    | Park et al. .         |         |

#### FOREIGN PATENT DOCUMENTS

WO 97/27  
884 A1 8/1997 (WO) .

#### OTHER PUBLICATIONS

K. Park, "Enzyme-digestible swelling hydrogels as platforms for long-term oral drug delivery: synthesis and characterization", *Biomaterials* (Sep. 1988), pp. 435-441, vol. 9, Butterworth & Co.

C.J. Benning, *Plastic Foams: the physics and chemistry of product performance and process technology*, vol. 1, Polymer Engineering & Technology, (1969) Chptrs. 6, 7, 9, 10, 14. Wiley-Interscience.

F. Rodriguez, *Principles of Polymer Systems*, 2<sup>nd</sup> ed. (1982) pp. 363-378, Hemisphere Publ. Corp.

Plastic Foams, Part II. K. Frisch and J. Saunders, ed. (1973) Chptrs. 12, 15, 17. Marcel Dekker, Inc.

F.A. Shutov, *Integral/Structural Polymer Foams: Technology, Properties and Applications*, (1986) Chptrs. 1, 21. Springer Verlag.

"Absorbent PVA Material Finds Medical Applications," Medical Product Mfg. News Hotline (Apr. 1, 1995).

Vichterle et al., Hydrophilic gels for biological use, *Nature*, 185: 117-118, 1960.

Shalaby et al., In vitro and in vivo studies of enzyme-digestible hydrogels for oral drug delivery, *J. Controlled. Rel.*, 19: 131-144, 1992A.

Shalaby et al., Use of ultrasound imaging and fluoroscopic imaging to study gastric retention of enzyme-digestible hydrogels, *Biomaterials*, 13:289-296, 1992B.

Chirila et al., Poly (2-hydroxyethyl methacrylate) sponges as implant materials: In vivo and in vitro evaluation of cellular invasion, *Biomaterials*, 14: 26-38, 1993.

Skelly et al., Novel macroporous hydrogel adsorbents for artificial liver support haemoperfusion systems, *Polymer*, 20: 1051-1052, 1979.

Oxley et al., Macroporous hydrogels for biomedical applications: Methodology and morphology, *Biomaterials*, 14: 26-38, 1993.

Barvic et al., Biologic properties and possible uses of polymer-like sponges, *J. Biomed. Mater. Res.*, 1: 313-323, 1967.

(List continued on next page.)

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(57)

#### ABSTRACT

A superporous hydrogel composite is formed by polymerizing one or more ethylenically-unsaturated monomers, and a multiolefinic crosslinking agent, in the presence of particles of a disintegrant and a blowing agent. The disintegrant, which rapidly absorbs water, serves to greatly increase the mechanical strength of the superporous hydrogel and significantly shorten the time required to absorb water and swell. Superporous hydrogel composites prepared by this method have an average pore size in the range of 10  $\mu$ m to 3,000  $\mu$ m. Preferred particles of disintegrant include natural and synthetic charged polymers, such as crosslinked sodium carboxymethylcellulose, crosslinked sodium starch glycolate, and crosslinked polyvinylpyrrolidone. The blowing agent is preferably a compound that releases gas bubbles upon acidification, such as NaHCO<sub>3</sub>. Improved hydrogel composites formed without a blowing agent are also provided.

40 Claims, 5 Drawing Sheets